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FORM PTO-1390 (REV 5-93)

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## U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE

ATTORNEY DOCKET NO. 026220-00082

DATE: May 1, 2006

TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US) CONCERNING A FILING UNDER 35 U.S.C. 371

INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE PRIORITY DATE CLAIMED PCT/EP2004/013683 December 1, 2004 December 2, 2003 TITLE OF INVENTION: NITROOXYDERIVATIVES OF CARVEDILOL AND OTHER BETA BLOCKERS AS ANTIHYPERTENSIVE **DRUGS** APPLICANT(S) FOR DO/EO/US: Piero DEL SOLDATO (Milano, Italy), Francesca BENEDINI (Milano, Italy) and Ennio ONGINI (Milano, Italy) Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: This is a FIRST submission of items concerning a filing under 35 U.S.C. 371. This is a SECOND or SUBSEQUENT submission of items concerning a filing under 35 U.S.C. 371. This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below. The U.S. has been elected (Article 31). A copy of the International Application as filed [35 U.S.C. 371(c)(2)] a. is attached hereto (required only if not transmitted by the International Bureau). b. A has been communicated by the International Bureau. c. is not required, as the application was filed in the United States Receiving Office (RO/US). 6. An English language translation of the International Application into English [35 U.S.C. 371(c)(2)]. a. is attached hereto. b. has been previously submitted under 35 U.S.C. 154(d)(4). 7. Amendments to the claims of the International Application under PCT Article 19 [35 U.S.C. 371(c)(3)] a. are attached hereto (required only if not transmitted by the International Bureau). b. have been communicated by the International Bureau. c. have not been made; however, the time limit for making such amendments has NOT expired. d. have not been made and will not be made. An English language translation of the amendments to the claims under PCT Article 19 [35 U.S.C. 371(c)(3)]. 9. An oath or declaration of the inventor(s) [35 U.S.C. 371(c)(4)]. Article 36 [35 U.S.C. 371(c)(5)]. Items 11 to 20 below concern other document(s) or information included: 11. An Information Disclosure Statement under 37 C.F.R. 1.97 and 1.98; and PTO-SB08a Form. 12. An assignment document for recording. A separate cover sheet in compliance with 37 C.F.R. 3.28 and 3.31 is included. A preliminary amendment. 14. An Application Data Sheet under 37 CFR 1.76. 15. A substitute specification. 16. A power of attorney and/or change of address letter. 17. A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821 - 1.825. 18. A second copy of the published international application under 35 U.S.C. 154(d)(4). 19. A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4). Other items or information: International Search Report, Form PCT/IB/304, Form PCT/IB/308 (First Notice), Form PCT/IB/301, And Form PCT/ISA/237

IAP12 Rec'd PCT/FTO 01 MAY 2006

| U.S. APPLN. NO. (IF KNOWN, O. 1) INTERNATIONAL APPLICATION NO.  |              |           |   |                                       |            |            | ATTORNEY DOCKET NO.                       |                    |              |
|---|--------------|-----------|---|---------------------------------------|------------|------------|---|--------------------|--------------|
| SEE 37 C.F.R. 1.50 / 9 1 PCT/EP2004/013683  The following fees are submitted:   |              |           |   |                                       |            |            | 026220-00082  CALCULATIONS   PTO USE ONLY |                    |              |
| 21. Basic National Fee \$300.00   |              |           |   |                                       |            |            | S.  | 300.00             | FIGUSEONLI   |
| 22. Examination Fee  If International Preliminary Examination Report prepared by USPTO  and all claims satisfy provisions of PCT Article 33(1)-(4) \$100.00  All other situations \$200.00  |              |           |   |                                       |            |            | \$  | 200.00             |              |
| 23. Search Fee Search Fee (37 CFR 1.445(a)(2)) has been paid on the International Application to the USPTO as an International Searching Authority International Search Report prepared and provided to the Office \$400.00 All other situations \$500.00   |              |           |   |                                       |            |            | \$  | 400.00             |              |
| TOTAL OF CALCULATIONS FOR 21, 22 and 23 =   |              |           |   |                                       |            |            | \$  | 900.00             |              |
| Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing or computer program listing filed in an electronic medium). The fee is \$250 for each additional 50 sheets of paper or fraction thereof.   |              |           |   |                                       |            |            |   |                    |              |
| Total Sheets  | Extra Sheets |           | Number of each additional 50 of thereof (round up to a whole nu |                                       |            | Rate       |   |                    |              |
| - 100 =   |              | /50 =     |   |                                       |            | x \$250.00 | \$  | 0.00               |              |
| Surcharge of \$130.00 for furnishing the oath or declaration later than 20 30 months from the earliest claimed priority date [37 C.F.R. 1.492(e)].  |              |           |   |                                       |            |            | \$  | 0.00               | <del>.</del> |
| Claims  |              |           | per Filed Number Extra  |                                       | Rate       |            |   |                    |              |
| Total Claims  |              | 98 - 20 = |   | 78                                    | x \$ 50.00 |            | \$  | 3900.00            |              |
| Independent Claims 1 - 3 =  |              |           | =   | 0 x \$200.00                          |            |            | \$  | 0.00               |              |
| Multiple dependent claim(s) (if applicable)   |              |           |   |                                       | + \$360.00 |            |   | 360.00             |              |
| TOTAL OF ABOVE CALCULATIONS =   |              |           |   |                                       |            |            | \$  | 5160.00            |              |
| Reduction by one-half for filing by small entity, if applicable. Small entity status under 37 C.F.R. §§1.9 and 1.27 is hereby claimed.  |              |           |   |                                       |            |            | \$  |                    |              |
| SUBTOTAL =  |              |           |   |                                       |            |            | \$  | 5160.00            |              |
| Processing fee of \$130.00 for furnishing the English translation later the 20 30 months from the earliest claimed priority date [37 C.F.R. 1.492(f)].  |              |           |   |                                       |            |            | \$  |                    |              |
| TOTAL NATIONAL FEE =  |              |           |   |                                       |            |            | \$  | 5160.00            |              |
| Fee for recording the enclosed assignment [37 C.F.R. 1.21(h)]. The assignment must be accompanied by an appropriate cover sheet (37 C.F.R. 3.28, 3.31). \$40.00 per property +  |              |           |   |                                       |            |            | \$  | 40.00              |              |
| TOTAL FEES ENCLOSED =   |              |           |   |                                       |            |            | \$  | 5200.00            |              |
|   |              |           |   |                                       |            |            |   | ount to be<br>nded | \$           |
|   |              |           |   |                                       |            |            | Cha                                       | rged               | \$           |
| <ul> <li>a. Check # in the amount of \$\\$ to cover the above fees is enclosed.</li> <li>b. Please charge my Deposit Account No. 01-2300 in the amount of \$\\$ \\$5200.00 to cover the above fee.         A duplicate copy of this sheet is enclosed.     </li> <li>c. Check # in the amount of \$\\$ \\$5200.00 to cover the above fee.</li> <li>A duplicate copy of this sheet is enclosed.</li> <li>The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 01-2300.</li> </ul> |              |           |   |                                       |            |            |   |                    |              |
| NOTE: Where an appropriate time limit under 37 C.F.R. 1.494 or 1.495 has not been met, a petition to revive (37 C.F.R. 1.137(a) or (b)) must be filed and granted to restore the application to pending status.   |              |           |   |                                       |            |            |   |                    |              |
| SEND ALL CORRESPONDENCE TO: Customer No. 004372   |              |           |   |                                       |            |            |   |                    |              |
| Arent Fox PLLC 1050 Connecticut Avenue, N.W.  Richard J. Berman   |              |           |   |                                       |            |            |   |                    | <i>)</i>     |
| Suite 400  Reg. No. 39,107  Washington, D.C. 20036-5339   |              |           |   |                                       |            |            |   |                    |              |
| Tel: (202) 857-600  |              |           | 310   | The PTO did not receive the following |            |            |   | RJI                | 3/cdw        |
|   |              |           |   | listed item(s)_                       | NO940      | 100 15     |   |                    |              |